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LOGINID:SSSPTA1642BJF

PASSWORD:
TERMINAL (ENTER 1, 2, 3, OR ?):2

* * * * * * * * * * Welcome to STN International * * * * * * * * * *

NEWS 1		Web Page for STN Seminar Schedule - N. America
NEWS 2	JAN 02	STN pricing information for 2008 now available
NEWS 3	JAN 16	CAS patent coverage enhanced to include exemplified prophetic substances
NEWS 4	JAN 28	USPATFULL, USPAT2, and USPATOLD enhanced with new custom IPC display formats
NEWS 5	JAN 28	MARPAT searching enhanced
NEWS 6	JAN 28	USGENE now provides USPTO sequence data within 3 days of publication
NEWS 7	JAN 28	TOXCENTER enhanced with reloaded MEDLINE segment
NEWS 8	JAN 28	MEDLINE and LMEDLINE reloaded with enhancements
NEWS 9	FEB 08	STN Express, Version 8.3, now available
NEWS 10	FEB 20	PCI now available as a replacement to DPCI
NEWS 11	FEB 25	IFIREF reloaded with enhancements
NEWS 12	FEB 25	IMSPRODUCT reloaded with enhancements
NEWS 13	FEB 29	WPINDEX/WPIDS/WPIX enhanced with ECLA and current U.S. National Patent Classification
NEWS 14	MAR 31	IFICDB, IFIPAT, and IFIUDB enhanced with new custom IPC display formats
NEWS 15	MAR 31	CAS REGISTRY enhanced with additional experimental spectra
NEWS 16	MAR 31	CA/CAplus and CASREACT patent number format for U.S. applications updated
NEWS 17	MAR 31	LPCI now available as a replacement to LDPCI
NEWS 18	MAR 31	EMBASE, EMBAL, and LEMBASE reloaded with enhancements
NEWS 19	APR 04	STN AnaVist, Version 1, to be discontinued
NEWS 20	APR 15	WPIDS, WPINDEX, and WPIX enhanced with new predefined hit display formats
NEWS 21	APR 28	EMBASE Controlled Term thesaurus enhanced
NEWS 22	APR 28	IMSRESEARCH reloaded with enhancements
NEWS 23	MAY 30	INPAFAMDB now available on STN for patent family searching
NEWS 24	MAY 30	DGENE, PCTGEN, and USGENE enhanced with new homology sequence search option
NEWS 25	JUN 06	EPFULL enhanced with 260,000 English abstracts
NEWS 26	JUN 06	KOREAPAT updated with 41,000 documents
NEWS 27	JUN 13	USPATFULL and USPAT2 updated with 11-character patent numbers for U.S. applications
NEWS 28	JUN 19	CAS REGISTRY includes selected substances from web-based collections
NEWS 29	JUN 25	CA/CAplus and USPAT databases updated with IPC reclassification data
NEWS 30	JUN 30	AEROSPACE enhanced with more than 1 million U.S. patent records
NEWS 31	JUN 30	EMBASE, EMBAL, and LEMBASE updated with additional options to display authors and affiliated

```
                  organizations
 NEWS 32  JUN 30  STN on the Web enhanced with new STN AnaVist
                  Assistant and BLAST plug-in
 NEWS 33  JUN 30  STN AnaVist enhanced with database content from EPFULL

 NEWS EXPRESS JUNE 27 08 CURRENT WINDOWS VERSION IS V8.3,
              AND CURRENT DISCOVER FILE IS DATED 23 JUNE 2008.

 NEWS HOURS    STN Operating Hours Plus Help Desk Availability
 NEWS LOGIN    Welcome Banner and News Items
 NEWS IPC8     For general information regarding STN implementation of IPC 8

Enter NEWS followed by the item number or name to see news on that
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  research.  Use for software development or design or implementation
  of commercial gateways or other similar uses is prohibited and may
  result in loss of user privileges and other penalties.

* * * * * * * * * * * * * * * * STN  Columbus  * * * * * * * * * * * * * * * *

FILE 'HOME' ENTERED AT 08:59:51 ON 07 JUL 2008

=> file reg
COST IN U.S. DOLLARS                                SINCE FILE      TOTAL
                                                         ENTRY    SESSION
FULL ESTIMATED COST                                       0.21       0.21

FILE 'REGISTRY' ENTERED AT 09:00:16 ON 07 JUL 2008
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
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Property values tagged with IC are from the ZIC/VINITI data file
provided by InfoChem.

STRUCTURE FILE UPDATES:    6 JUL 2008  HIGHEST RN 1032827-24-9
DICTIONARY FILE UPDATES:   6 JUL 2008  HIGHEST RN 1032827-24-9

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 9, 2008.

  Please note that search-term pricing does apply when
  conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and
predicted properties as well as tags indicating availability of
experimental property data in the original document.  For information
on property searching in REGISTRY, refer to:

http://www.cas.org/support/stngen/stndoc/properties.html

=> E "EPOTHILONE B"/CN 25
E1           1     EPOTHILONE A8/CN
E2           1     EPOTHILONE A9/CN
E3           1 --> EPOTHILONE B/CN
E4           1     EPOTHILONE B (12R,13R) ACETONIDE/CN
E5           1     EPOTHILONE B A-EPOXIDE/CN
E6           1     EPOTHILONE B ACID/CN
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E7            1     EPOTHILONE B HYDROXYLASE/CN
E8            1     EPOTHILONE B HYDROXYLASE (AMYCOLATOPSIS ORIENTALIS GENE EBH)/CN
E9            1     EPOTHILONE B N-OXIDE/CN
E10           1     EPOTHILONE B10/CN
E11           1     EPOTHILONE C/CN
E12           1     EPOTHILONE C BIS(TERT-BUTYLDIMETHYLSILYL) ETHER/CN
E13           1     EPOTHILONE C/D 12,13-EPOXIDASE/CN
E14           1     EPOTHILONE C/D MONOOXYGENASE/CN
E15           1     EPOTHILONE C/D SYNTHETASE/CN
E16           1     EPOTHILONE C1/CN
E17           1     EPOTHILONE C2/CN
E18           1     EPOTHILONE C3/CN
E19           1     EPOTHILONE C4/CN
E20           1     EPOTHILONE C5/CN
E21           1     EPOTHILONE C6/CN
E22           1     EPOTHILONE C7/CN
E23           1     EPOTHILONE C8/CN
E24           1     EPOTHILONE C9/CN
E25           1     EPOTHILONE D/CN

=> S E3
L1            1 "EPOTHILONE B"/CN

=> S L1 EXA SAM
SAMPLE IS IGNORED AS A SCOPE FOR THIS SEARCH
L2            1 "EPOTHILONE B"/CN

=> DIS L2 1 SAM
THE ESTIMATED COST FOR THIS REQUEST IS 1.04 U.S. DOLLARS
DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N:Y

L2   ANSWER 1 OF 1  REGISTRY  COPYRIGHT 2008 ACS on STN
IN   4,17-Dioxabicyclo[14.1.0]heptadecane-5,9-dione, 7,11-dihydroxy-
     8,8,10,12,16-pentamethyl-3-[(1E)-1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-
     , (1S,3S,7S,10R,11S,12S,16R)-
MF   C27 H41 N O6 S

Absolute stereochemistry.  Rotation (-).
Double bond geometry as shown.
```

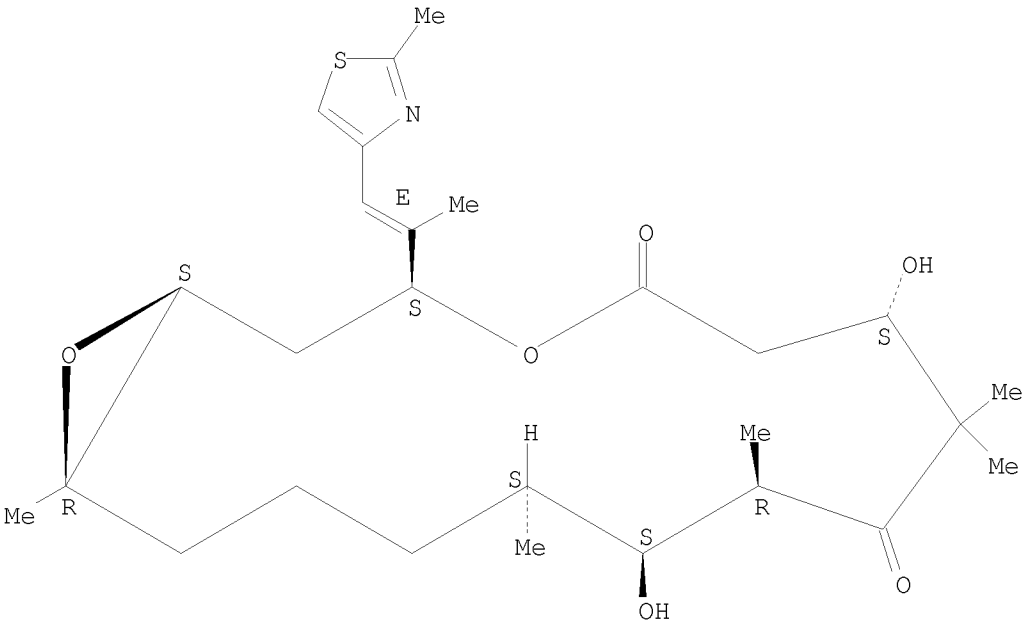

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

=>

---Logging off of STN---

=>
Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	13.64	13.85

STN INTERNATIONAL LOGOFF AT 09:03:08 ON 07 JUL 2008

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Welcome to STN International! Enter x:x

LOGINID:SSSPTA1642BJF

PASSWORD:
TERMINAL (ENTER 1, 2, 3, OR ?):2

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* * * * * * * * * *   Welcome to STN International   * * * * * * * * * *

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                   prophetic substances
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                   custom IPC display formats
 NEWS  5  JAN 28   MARPAT searching enhanced
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 NEWS 24  MAY 30   DGENE, PCTGEN, and USGENE enhanced with new homology
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 NEWS 25  JUN 06   EPFULL enhanced with 260,000 English abstracts
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                   patent numbers for U.S. applications
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                   web-based collections
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 NEWS 30  JUN 30   AEROSPACE enhanced with more than 1 million U.S.
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 NEWS 31  JUN 30   EMBASE, EMBAL, and LEMBASE updated with additional
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                   organizations
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* * * * * * * * * * * * * * * * STN Columbus * * * * * * * * * * * * * * * *

FILE 'HOME' ENTERED AT 10:09:50 ON 07 JUL 2008

=> file pctfull
COST IN U.S. DOLLARS   SINCE FILE   TOTAL
                                                         ENTRY   SESSION
FULL ESTIMATED COST   0.21   0.21

FILE 'PCTFULL' ENTERED AT 10:10:27 ON 07 JUL 2008
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FILE LAST UPDATED: 4 JUL 2008   <20080704/UP>
FILE COVERS 1978 TO DATE

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>>> NEW FIELD UPTX, FIELD /EW NO LONGER AVAILBLE - SEE HELP CHANGE <<<

=> s epothilon?
L1   2484 EPOTHILON?

=> s l1/ab or l1/ti
           144 EPOTHILON?/AB
           129 EPOTHILON?/TI
L2   159 (EPOTHILON?/AB) OR (EPOTHILON?/TI)

=> s l2 not py>2001
        817323 PY>2001
L3   53 L2 NOT PY>2001

=> s combination and l3
        567168 COMBINATION
        264042 COMBINATIONS
        617900 COMBINATION
                  (COMBINATION OR COMBINATIONS)
L4   33 COMBINATION AND L3

=> d ibib 1-5

L4   ANSWER 1 OF 33   PCTFULL   COPYRIGHT 2008 Univentio on STN
ACCESSION NUMBER: 2001092255 PCTFULL ED 20020826
TITLE (ENGLISH):   EPOTHILONE DERIVATIVES AND METHODS FOR MAKING
                        AND USING THE SAME
TITLE (FRENCH):   DERIVES D'EPOTHILONE, PROCEDES DE PRODUCTION
                        ET METHODES D'UTILISATION
INVENTOR(S):   SANTI, Daniel;
                        FARDIS, Maria;
                        ASHLEY, Gary
PATENT ASSIGNEE(S):   KOSAN BIOSCIENCES, INC.;
                        SANTI, Daniel;
                        FARDIS, Maria;
                        ASHLEY, Gary
DOCUMENT TYPE: Patent

PATENT INFORMATION:

NUMBER	KIND	DATE
WO 2001092255	A2	20011206

DESIGNATED STATES

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR
CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID
IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD
MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL
TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW GH GM KE LS MW
MZ SD SL SZ TZ UG ZW AM AZ BY KG KZ MD RU TJ TM AT BE
CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE TR BF
BJ CF CG CI CM GA GN GW ML MR NE SN TD TG

PRIORITY INFO.: US 2000-60/207,655 20000526
US 2000-60/218,260 20000714
US 2000-60/231,552 20000911

APPLICATION INFO.: WO 2001-US15763 A 20010515

L4 ANSWER 2 OF 33 PCTFULL COPYRIGHT 2008 Univentio on STN

ACCESSION NUMBER: 2001083800 PCTFULL ED 20020826

TITLE (ENGLISH): PRODUCTION OF POLYKETIDES

TITLE (FRENCH): PRODUCTION DE POLYKETIDES

INVENTOR(S): ARSLANIAN, Robert, L.;
ASHLEY, Gary;
FRYKMAN, Scott;
JULIEN, Bryan;
KATZ, Leonard;
KHOSLA, Chaitan;
LAU, Janice;
LICARDI, Peter, J.;
REGENTIN, Rika;
SANTI, Daniel;
TANG, Li

PATENT ASSIGNEE(S): KOSAN BIOSCIENCES, INC.;
ARSLANIAN, Robert, L.;
ASHLEY, Gary;
FRYKMAN, Scott;
JULIEN, Bryan;
KATZ, Leonard;
KHOSLA, Chaitan;
LAU, Janice;
LICARDI, Peter, J.;
REGENTIN, Rika;
SANTI, Daniel;
TANG, Li

DOCUMENT TYPE: Patent

PATENT INFORMATION:

NUMBER	KIND	DATE
WO 2001083800	A2	20011108

DESIGNATED STATES

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU
CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN
IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK
MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM
TR TT TZ UA UG US UZ VN YU ZA ZW GH GM KE LS MW MZ SD
SL SZ TZ UG ZW AM AZ BY KG KZ MD RU TJ TM AT BE CH CY
DE DK ES FI FR GB GR IE IT LU MC NL PT SE TR BF BJ CF
CG CI CM GA GN GW ML MR NE SN TD TG

PRIORITY INFO.: US 2000-09/560,367 20000428
US 2000-60/232,696 20000914
US 2000-60/257,517 20001221

	US 2001-09/825,856		20010403
	US 2001-09/825,876		20010403
	US 2001-60/269,020		20010413
APPLICATION INFO.:	WO 2001-US13793	A	20010426

L4 ANSWER 3 OF 33 PCTFULL COPYRIGHT 2008 Univentio on STN
ACCESSION NUMBER: 2001081341 PCTFULL ED 20020826
TITLE (ENGLISH): 9-OXA-EPOTHILON DERIVATIVES, METHOD FOR THE PRODUCTION AND USE THEREOF IN PHARMACEUTICAL PREPARATIONS
TITLE (FRENCH): DERIVES DE 9-OXA-EPOTHILONE, LEUR PROCEDE DE PRODUCTION ET LEUR UTILISATION PHARMACEUTIQUE
INVENTOR(S): SCHWEDE, Wolfgang;
KLAR, Ulrich;
SKUBALLA, Werner;
BUCHMANN, Bernd;
HOFFMANN, Jens;
LICHTNER, Rosemarie
PATENT ASSIGNEE(S): SCHERING AKTIENGESELLSCHAFT;
SCHWEDE, Wolfgang;
KLAR, Ulrich;
SKUBALLA, Werner;
BUCHMANN, Bernd;
HOFFMANN, Jens;
LICHTNER, Rosemarie
DOCUMENT TYPE: Patent
PATENT INFORMATION:

NUMBER	KIND	DATE
WO 2001081341	A2	20011101

DESIGNATED STATES
W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW GH GM KE LS MW MZ SD SL SZ TZ UG ZW AM AZ BY KG KZ MD RU TJ TM AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE TR BF BJ CF CG CI CM GA GN GW ML MR NE SN TD TG
PRIORITY INFO.: DE 2000-100 20 899.1 20000420
APPLICATION INFO.: WO 2001-EP4551 A 20010419

L4 ANSWER 4 OF 33 PCTFULL COPYRIGHT 2008 Univentio on STN
ACCESSION NUMBER: 2001073103 PCTFULL ED 20020822
TITLE (ENGLISH): PREPARATION OF EPOTHILONE INTERMEDIATES
TITLE (FRENCH): PREPARATION D'INTERMEDIAIRES D'EPOTHILONE
INVENTOR(S): VITE, Gregory, D.;
KIM, Soong-Hoon;
HOeEFLE, Gerhard
PATENT ASSIGNEE(S): BRISTOL-MYERS SQUIBB COMPANY;
VITE, Gregory, D.;
KIM, Soong-Hoon;
HOeEFLE, Gerhard
DOCUMENT TYPE: Patent
PATENT INFORMATION:

NUMBER	KIND	DATE
WO 2001073103	A2	20011004

DESIGNATED STATES
W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG

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                        MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ
                        TM TR TT TZ UA UG US UZ VN YU ZA ZW GH GM KE LS MW MZ
                        SD SL SZ TZ UG ZW AM AZ BY KG KZ MD RU TJ TM AT BE CH
                        CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE TR BF BJ
                        CF CG CI CM GA GN GW ML MR NE SN TD TG
PRIORITY INFO.:         US 2000-60/191,975      20000324
APPLICATION INFO.:      WO 2001-US9620       A  20010323

L4     ANSWER 5 OF 33    PCTFULL   COPYRIGHT 2008 Univentio on STN
ACCESSION NUMBER:       2001070716 PCTFULL  ED 20020822
TITLE (ENGLISH):        A PROCESS FOR THE PREPARATION OF EPOTHILONE
                        ANALOGS AND INTERMEDIATES
TITLE (FRENCH):         PREPARATION D'ANALOGUES ET D'INTERMEDIAIRES D'
                        EPOTHILONE
INVENTOR(S):            LI, Wen, Sen;
                        THORNTON, John, E.;
                        GUO, Zhenrong;
                        SWAMINATHAN, Shankar;
                        MCCONLOGUE, Gary, W.
PATENT ASSIGNEE(S):     BRISTOL-MYERS SQUIBB COMPANY;
                        LI, Wen, Sen;
                        THORNTON, John, E.;
                        GUO, Zhenrong;
                        SWAMINATHAN, Shankar;
                        MCCONLOGUE, Gary, W.
DOCUMENT TYPE:          Patent
PATENT INFORMATION:
                        NUMBER             KIND     DATE
                        --------------------------------
                        WO 2001070716        A1 20010927
DESIGNATED STATES
       W:               AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR
                        CU CZ DE DK DM EE ES FI GB GD GE GH GM HR HU ID IL IN
                        IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK
                        MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM
                        TR TT TZ UA UG US UZ VN YU ZA ZW GH GM KE LS MW MZ SD
                        SL SZ TZ UG ZW AM AZ BY KG KZ MD RU TJ TM AT BE CH CY
                        DE DK ES FI FR GB GR IE IT LU MC NL PT SE TR BF BJ CF
                        CG CI CM GA GN GW ML MR NE SN TD TG
PRIORITY INFO.:         US 2000-09/528,526      20000320
APPLICATION INFO.:      WO 2001-US7749       A  20010312


=> d his

     (FILE 'HOME' ENTERED AT 10:09:50 ON 07 JUL 2008)

     FILE 'PCTFULL' ENTERED AT 10:10:27 ON 07 JUL 2008
L1         2484 S EPOTHILON?
L2          159 S L1/AB OR L1/TI
L3           53 S L2 NOT PY>2001
L4           33 S COMBINATION AND L3

=> s l4 and (taxol or paclitaxel)
           9622 TAXOL
            272 TAXOLS
           9705 TAXOL
                  (TAXOL OR TAXOLS)
          10390 PACLITAXEL
             72 PACLITAXELS
          10392 PACLITAXEL
                  (PACLITAXEL OR PACLITAXELS)
```

```
L5          29 L4 AND (TAXOL OR PACLITAXEL)

=> s l5 and Her?
        988529 HER?
L6          29 L5 AND HER?

=> s l5 and (HER2 or HER-2)
          4722 HER2
        118696 HER
          1043 HERS
        119313 HER
                  (HER OR HERS)
       1276185 2
          3260 HER-2
                  (HER(W)2)
L7           1 L5 AND (HER2 OR HER-2)

=> d ibib abs

L7     ANSWER 1 OF 1     PCTFULL   COPYRIGHT 2008 Univentio on STN
ACCESSION NUMBER:       1999002514 PCTFULL  ED 20020515
TITLE (ENGLISH):        EPOTHILONE DERIVATIVES
TITLE (FRENCH):         DERIVES D'EPOTHILONE
INVENTOR(S):            VITE, Gregory, D.;
                        BORZILLERI, Robert, M.;
                        KIM, Soong-Hoon;
                        JOHNSON, James, A.
PATENT ASSIGNEE(S):     BRISTOL-MYERS SQUIBB COMPANY
LANGUAGE OF PUBL.:      English
DOCUMENT TYPE:          Patent
PATENT INFORMATION:
                        NUMBER             KIND     DATE
                        --------------------------------
                        WO 9902514           A2 19990121
DESIGNATED STATES
       W:               AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE
                        ES FI GB GE GH GM GW HU ID IL IS JP KE KG KP KR KZ LC
                        LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU
                        SD SE SG SI SK SL TJ TM TR TT UA UG UZ VN YU ZW GH GM
                        KE LS MW SD SZ UG ZW AM AZ BY KG KZ MD RU TJ TM AT BE
                        CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE BF BJ
                        CF CG CI CM GA GN ML MR NE SN TD TG
PRIORITY INFO.:         US 1997-60/051,951      19970708
                        US 1997-60/067,524      19971204
APPLICATION INFO.:      WO 1998-US12550      A  19980616
ABEN   The present invention relates to compounds of formula (I), Q is selected
       from the group
       consisting of (II), G is selected from the group consisting of alkyl,
       substituted alkyl, substituted
       or or unsubstituted aryl, heterocyclo, (III), W is O or NR15; X is O or
       H,H; Y is selected from the
       group consisting of O; H,OR16; OR17,OR17; NOR18; H,NOR19; H,NR20R21;
       H,H; or CHR22; OR17OR17 can be
       a cyclic ketal; Z1 and Z2 are selected from the group consisting of CH2,
       O, NR23, S or SO2, wherein
       only one of Z and Z2 is a heteroatom; B1 and B2 are selected from the
       group consisting of OR24, or
       OCOR25, or 2CNR26R27; when B1 is H and Y is OH, H they can form a
       six-membered ring ketal or acetal;
       D is selected from the group consisting of NR28R29, NR30COR31 or
       saturated heterocycle R1, R2, R3,
       R4, R5, R6, R7, R13, R14, R18, R19, R20, R21, R22, R26 and R27 are
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selected from the group H, alkyl,
substituted alkyl, or aryl and when R1 and R2 are alkyl can be joined to form a cycloalkyl; R3 and
R4 are alkyl can be joined to form a cycloalkyl; R9, R10, R16, R17, R24, R25, and R31 are selected
from the group H, alkyl, or substituted alkyl; R8, R11, R12, R28, R30, R32, R33, and R30 are
selected from the group consisting of H, alkyl, substituted alkyl, aryl, substituted aryl,
cycloalkyl, or heterocyclo; R15, R23 and R29 are selected from the group consisting of H, alkyl,
substituted alkyl, aryl, substituted aryl, cycloalkyl, heterocyclo, R32C=O, R33SO2, hydroxy, O-alkyl
or O-substituted alkyl, the pharmaceutically acceptable salts thereof and any hydrates, solvates or
geometric, optical and stereoisomers thereof, with the proviso that compounds wherein: W and X are
both O; and R1, R2, R7 are H; and R3, R4, R6, are methyl; and R8, is H or methyl; and Z1, and Z2,
are CH2; and G is 1-methyl-2-(substituted-4-thiazolyl)ethenyl; and Q is as defined above are
excluded.

ABFR La presente invention concerne des composes de la formule (I) dans laquelle Q est selectionne
dans le groupe constitue par le groupement (II); G est selectionne dans le groupe constitue par
alkyle, akyle substitue, aryle substitue ou insusbstitue, heterocyclo, le groupement (III); W est O
ou NR15; X est O ou H,H; Y est selectionne dans le groupe constitue par O; H,OR16; OR17,OR17; NOR18;
H,NOR19; H,NR20R21; H,H; ou CHR22; OR17,OR17 pouvant etre un cetal cyclique; Z1 et Z2 sont
selectionnes dans le groupe constitue par CH2, O, NR23, S ou SO2, dans lequel seuls Z et Z2 sont un
heteroatome; B1 et B2 sont selectionnes dans le groupe constitue par OR24 ou OCOR25 ou O2CNR26R27;
et peuvent former ensemble un noyau cetal ou acetal a six chainons si B1 est H et Y est OH,H; D est
selectionne dans le groupe constitue par NR28R29, NR30COR31 ou un heterocycle sature. R1, R2, R3,
R4, R5, R6, R7, R13, R14, R18, R19, R20, R21, R22, R26 et R27 sont selectionnes dans le groupe
constitue par H, alkyle, alkyle substitue ou aryle, et peuvent former ensemble un cycloalkyle si R1
et R2 ou R3 et R4 sont alkyle; R9, R10, R16, R17, R24, R25 et R31 sont selectionnes dans le groupe
constitue par H, alkyle ou alkyle substitue; R8, R11, R12, R28, R30, R32, R33 et R30 sont
selectionnes dans le groupe constitue par H, alkyle, alkyle substitue, aryle, aryle substitue,
cycloalkyle ou heterocyclo; R15, R23 et R29 sont selectionnes dans le groupe constitue par H,
alkyle, alkyle substitue, aryle, aryle substitue, cycloalkyle ou heterocyclo, R32C=O, R33SO2,
hydroxy, O-alkyle ou O-alkyle substitue, leurs sels pharmaceutiquement acceptables ou leurs
eventuels hydrates, solvates ou isomeres geometriques, optiques, ou stereoisomeres, a condition que
soient exclus les composes dans lesquels W et X sont tous deux O; et R1, R2 et R7 sont H; et R3, R4
et R6 sont methyle; et R8 est H ou methyle; et Z1 et Z2 sont CH2; et G est

1-methyl-2-(substitue-4-thiazolyl)ethenyle; et Q est tel que defini ci-dessus.

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=> d his

     (FILE 'HOME' ENTERED AT 10:09:50 ON 07 JUL 2008)

     FILE 'PCTFULL' ENTERED AT 10:10:27 ON 07 JUL 2008
L1          2484 S EPOTHILON?
L2           159 S L1/AB OR L1/TI
L3            53 S L2 NOT PY>2001
L4            33 S COMBINATION AND L3
L5            29 S L4 AND (TAXOL OR PACLITAXEL)
L6            29 S L5 AND HER?
L7             1 S L5 AND (HER2 OR HER-2)

=> s l6 and (HER2 or HER-2)
          4722 HER2
        118696 HER
          1043 HERS
        119313 HER
                 (HER OR HERS)
       1276185 2
          3260 HER-2
                 (HER(W)2)
L8             1 L6 AND (HER2 OR HER-2)

=> s l5 and (HER2 or HER-2)
          4722 HER2
        118696 HER
          1043 HERS
        119313 HER
                 (HER OR HERS)
       1276185 2
          3260 HER-2
                 (HER(W)2)
L9             1 L5 AND (HER2 OR HER-2)

=> d ibib abs kwic

L9     ANSWER 1 OF 1     PCTFULL   COPYRIGHT 2008 Univentio on STN
ACCESSION NUMBER:       1999002514 PCTFULL  ED 20020515
TITLE (ENGLISH):        EPOTHILONE DERIVATIVES
TITLE (FRENCH):         DERIVES D'EPOTHILONE
INVENTOR(S):            VITE, Gregory, D.;
                        BORZILLERI, Robert, M.;
                        KIM, Soong-Hoon;
                        JOHNSON, James, A.
PATENT ASSIGNEE(S):     BRISTOL-MYERS SQUIBB COMPANY
LANGUAGE OF PUBL.:      English
DOCUMENT TYPE:          Patent
PATENT INFORMATION:
                        NUMBER              KIND     DATE
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                        WO 9902514            A2 19990121
DESIGNATED STATES
       W:               AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE
                        ES FI GB GE GH GM GW HU ID IL IS JP KE KG KP KR KZ LC
                        LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU
                        SD SE SG SI SK SL TJ TM TR TT UA UG UZ VN YU ZW GH GM
                        KE LS MW SD SZ UG ZW AM AZ BY KG KZ MD RU TJ TM AT BE
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CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE BF BJ
CF CG CI CM GA GN ML MR NE SN TD TG
PRIORITY INFO.: US 1997-60/051,951 19970708
US 1997-60/067,524 19971204
APPLICATION INFO.: WO 1998-US12550 A 19980616
ABEN The present invention relates to compounds of formula (I), Q is selected from the group consisting of (II), G is selected from the group consisting of alkyl, substituted alkyl, substituted or or unsubstituted aryl, heterocyclo, (III), W is O or NR15; X is O or H,H; Y is selected from the group consisting of O; H,OR16; OR17,OR17; NOR18; H,NOR19; H,NR20R21; H,H; or CHR22; OR17OR17 can be a cyclic ketal; Z1 and Z2 are selected from the group consisting of CH2, O, NR23, S or SO2, wherein only one of Z and Z2 is a heteroatom; B1 and B2 are selected from the group consisting of OR24, or OCOR25, or 2CNR26R27; when B1 is H and Y is OH, H they can form a six-membered ring ketal or acetal; D is selected from the group consisting of NR28R29, NR30COR31 or saturated heterocycle R1, R2, R3, R4, R5, R6, R7, R13, R14, R18, R19, R20, R21, R22, R26 and R27 are selected from the group H, alkyl, substituted alkyl, or aryl and when R1 and R2 are alkyl can be joined to form a cycloalkyl; R3 and R4 are alkyl can be joined to form a cycloalkyl; R9, R10, R16, R17, R24, R25, and R31 are selected from the group H, alkyl, or substituted alkyl; R8, R11, R12, R28, R30, R32, R33, and R30 are selected from the group consisting of H, alkyl, substituted alkyl, aryl, substituted aryl, cycloalkyl, or heterocyclo; R15, R23 and R29 are selected from the group consisting of H, alkyl, substituted alkyl, aryl, substituted aryl, cycloalkyl, heterocyclo, R32C=O, R33SO2, hydroxy, O-alkyl or O-substituted alkyl, the pharmaceutically acceptable salts thereof and any hydrates, solvates or geometric, optical and stereoisomers thereof, with the proviso that compounds wherein: W and X are both O; and R1, R2, R7 are H; and R3, R4, R6, are methyl; and R8, is H or methyl; and Z1, and Z2, are CH2; and G is 1-methyl-2-(substituted-4-thiazolyl)ethenyl; and Q is as defined above are excluded.

ABFR La presente invention concerne des composes de la formule (I) dans laquelle Q est selectionne dans le groupe constitue par le groupement (II); G est selectionne dans le groupe constitue par alkyle, akyle substitue, aryle substitue ou insusbstitue, heterocyclo, le groupement (III); W est O ou NR15; X est O ou H,H; Y est selectionne dans le groupe constitue par O; H,OR16; OR17,OR17; NOR18; H,NOR19; H,NR20R21; H,H; ou CHR22; OR17,OR17 pouvant etre un cetal cyclique; Z1 et Z2 sont selectionnes dans le groupe constitue par CH2, O, NR23, S ou SO2, dans lequel seuls Z et Z2 sont un heteroatome; B1 et B2 sont selectionnes dans le groupe constitue par OR24 ou OCOR25 ou O2CNR26R27; et peuvent former ensemble un noyau cetal ou acetal a six chainons si B1 est H et Y est OH,H; D est selectionne dans le groupe constitue par NR28R29, NR30COR31 ou un heterocycle sature. R1, R2, R3,

R4, R5, R6, R7, R13, R14, R18, R19, R20, R21, R22, R26 et R27 sont selectionnes dans le groupe
constitue par H, alkyle, alkyle substitue ou aryle, et peuvent former ensemble un cycloalkyle si R1
et R2 ou R3 et R4 sont alkyle; R9, R10, R16, R17, R24, R25 et R31 sont selectionnes dans le groupe
constitue par H, alkyle ou alkyle substitue; R8, R11, R12, R28, R30, R32, R33 et R30 sont
selectionnes dans le groupe constitue par H, alkyle, alkyle substitue, aryle, aryle substitue,
cycloalkyle ou heterocyclo; R15, R23 et R29 sont selectionnes dans le groupe constitue par H,
alkyle, alkyle substitue, aryle, aryle substitue, cycloalkyle ou heterocyclo, R32C=O, R33SO2,
hydroxy, O-alkyle ou O-alkyle substitue, leurs sels pharmaceutiquement acceptables ou leurs
eventuels hydrates, solvates ou isomeres geometriques, optiques, ou stereoisomeres, a condition que
soient exclus les composes dans lesquels W et X sont tous deux O; et R1, R2 et R7 sont H; et R3, R4
et R6 sont methyle; et R8 est H ou methyle; et Z1 et Z2 sont CH2; et G est
1-methyl-2-(substitue-4-thiazolyl)ethenyle; et Q est tel que defini ci-dessus.

TIEN EPOTHILONE DERIVATIVES

TIFR DERIVES D'EPOTHILONE

DETD R
S Me
j0H
N3],]'] '
0 Me
0 OH 0
I EpothiloneA R=H
II EpothiloneB R=Me
have been found to exert microtubule-stabilizing effects similar to TAXOL and hence cytotoxic activity against rapidly proliferating'cells,
such as, tumor cells or other hyperproliferative cellular disease, see .Angew. Chem. Int. Ed. Engl.,. . .

The compounds of this invention. are also useful in combination with known anti-cancer and cytotoxic agents and treatments, including radiation. If formulated as a fixed dose, such combination products
employ the compounds of this invention within the dosage range described below and the other pharmaceutically active agent within its approved dosage range. Compounds of formula V can be used sequentially with known anticancer or cytotoxic agents and treatment, including radiation when a combination formulation is inappropriate.

Especially useful are cytotoxic drug combinations wherein the second
drug chosen acts in a different phase of the cell cycle, e.g. S phase, than
the present compounds of. . .

. . .
Synthase Inhibitors,
DNA Cross Linking Agents
Topoisomerase I and II Inhibitors
DNA Alkylating Agents

Ribonucleoside Reductase Inhibitors
Cytotoxic Factors e.g. TNF-alpha or
Growth factor inhibitors e.g. HER 2 receptor MAB's
The present compounds may exist as multiple optical, geometric,
and stereoisomers. Included within the present invention are all such
isomers and. . .

. . .

potency is
accomplished following a modified procedure of Swindell, et al., (see
Swindell, C.S., Krauss, N.E., Horwitz, S.B., and Ringel, I. Biologically
active taxol analogues with deleted A-ring side chain
substituents and
variable C-2' configurations. J. Med. Chem. 34: 1176-1184, 1991). These
modifications, in part, result. . .

. . .

cells were incubated at 37' form 72 hours at which time the
tetrazolium dye, MTS at 333 gg/ml (final concentration), in
combination
with the electron coupling agent phenazine methosulfate at 25 gm (final
concentration) was added. A dehydrogenase enzyme in live cells
reduces the MTS. . .

=>

---Logging off of STN---

=>
Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	23.30	23.51

STN INTERNATIONAL LOGOFF AT 10:14:45 ON 07 JUL 2008